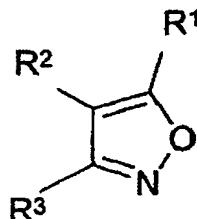


We claim

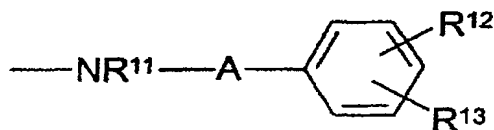
1. A substituted isoxazole derivative of the formula  
I



in which

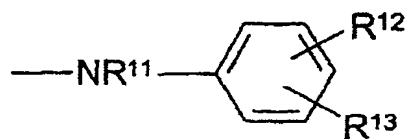
10 R<sup>1</sup> is selected from the group consisting of

- a) H;
- b) C<sub>1</sub>-C<sub>6</sub>-alkyl which may have 1 or 2 substituents independently of one another selected from the group consisting of NR<sup>4</sup>R<sup>5</sup> and OR<sup>6</sup>;
- 15 c) an aromatic or nonaromatic heterocycle having 5 or 6 ring atoms, including 1, 2 or 3 heteroatoms, independently of one another selected from the group consisting of N, O and S, where the heterocycle may have 1 or 2 substituents independently of one another selected from the group consisting of C<sub>1</sub>-C<sub>6</sub>-alkyl, halogen, CF<sub>3</sub>, OR<sup>6</sup>, NR<sup>7</sup>R<sup>8</sup>, NR<sup>9</sup>COR<sup>10</sup>, a radical of the formula II



25

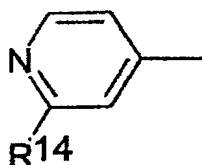
or a radical of the formula III



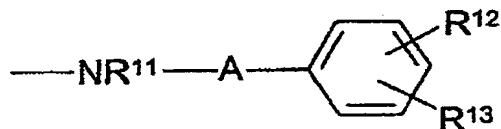
;

- d) phenyl which may have 1, 2 or 3 substituents independently of one another selected from the group consisting of  $\text{NR}^7\text{R}^8$ ,  $\text{OR}^6$ ,  $\text{C}_1\text{-C}_6\text{-alkyl}$ , halogen,  $\text{CF}_3$ ,  $\text{CN}$ ,  $\text{NO}_2$  and  $\text{CO}_2\text{R}^6$ ;
- 5 e) phenyl- $\text{C}_1\text{-C}_4\text{-alkyl}$ ;
- f)  $\text{C}_3\text{-C}_8\text{-cycloalkyl}$ ; and
- g)  $\text{NR}^7\text{R}^8$ ;

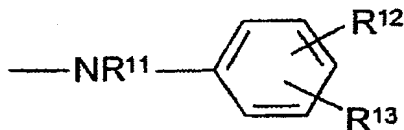
one of the radicals  $\text{R}^2$  and  $\text{R}^3$  is a radical of the  
10 formula IV



in which  $\text{R}^{14}$  is  $\text{C}_1\text{-C}_6\text{-alkyl}$ , halogen,  $\text{CF}_3$ ,  $\text{OR}^6$ ,  $\text{NR}^7\text{R}^8$ ,  
15  $\text{NR}^9\text{COR}^{10}$ , a radical of the formula

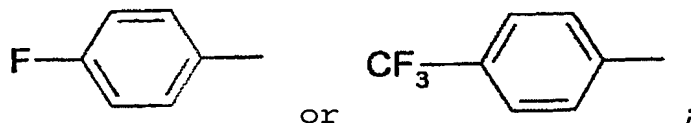


or a radical of the formula  
20



and

25 the second of the radicals  $\text{R}^2$  and  $\text{R}^3$  is



R<sup>4</sup> and R<sup>5</sup> independently of one another are H, C<sub>1</sub>-C<sub>6</sub>-alkyl, phenyl or phenyl-C<sub>1</sub>-C<sub>4</sub>-alkyl or together with the nitrogen atom to which they are attached form a saturated 5- or 6-membered heterocycle having 1 or 2  
5 heteroatoms independently of one another selected from the group consisting of N and O;

R<sup>6</sup>, R<sup>7</sup> and R<sup>8</sup> independently of one another are H or C<sub>1</sub>-C<sub>6</sub>-alkyl;

10 R<sup>9</sup> is H, C<sub>1</sub>-C<sub>6</sub>-alkyl or benzyl;

R<sup>10</sup> is C<sub>1</sub>-C<sub>6</sub>-alkyl, C<sub>3</sub>-C<sub>6</sub>-cycloalkyl or phenyl which may have 1 or 2 substituents independently of one another,  
15 selected from the group consisting of C<sub>1</sub>-C<sub>6</sub>-alkyl, C<sub>1</sub>-C<sub>6</sub>-alkoxy and halogen;

R<sup>11</sup> is H, C<sub>1</sub>-C<sub>6</sub>-alkyl or phenyl-C<sub>1</sub>-C<sub>4</sub>-alkyl;

20 R<sup>12</sup> and R<sup>13</sup> independently of one another are H, halogen, C<sub>1</sub>-C<sub>6</sub>-alkyl or C<sub>1</sub>-C<sub>6</sub>-alkoxy; and

A is straight-chain or branched C<sub>1</sub>-C<sub>6</sub>-alkylene; or  
25 an optical isomer or a physiologically acceptable salt thereof.

2. The compound as claimed in claim 1 where R<sup>2</sup> is 4-fluorophenyl and R<sup>3</sup> is the radical of the formula IV.

30 3. The compound as claimed in claim 1 where R<sup>2</sup> is the radical of the formula IV and R<sup>3</sup> is 4-fluorophenyl.

4. The compound as claimed in any of the preceding  
35 claims where R<sup>14</sup> is selected from the group consisting of halogen, OH, NR<sup>7</sup>R<sup>8</sup> and NR<sup>9</sup>COR<sup>10</sup>, where R<sup>7</sup> to R<sup>10</sup> have the meanings given in claim 1.

5. The compound as claimed in any of claims 1, 2 or 4 where  $R^1$  is H, phenyl which may have 1 or 2 halogen substituents,  $NR^7R^8$  or  $C_1-C_6$ -alkyl, where  $R^7$  and  $R^8$  have the meanings given in claim 1.

5

6. The compound as claimed in any of claims 1, 3 or 4, where  $R^1$  is  $C_1-C_6$ -alkyl which is substituted by  $NR^4R^5$  or  $OR^6$ , an aromatic heterocyclic radical having 5 or 6 ring atoms including 1 or 2 heteroatoms independently  
10 of one another selected from the group consisting of N and O, where the heterocycle is optionally substituted by  $NR^9COR^{10}$ , phenyl which is optionally substituted by  $NR^7R^8$  or  $C_1-C_6$ -alkoxy,  $NR^7R^8$  or  $C_3-C_6$ -cycloalkyl.

15 7. A pharmaceutical composition comprising at least one compound as claimed in any of claims 1 to 6, if appropriate together with one or more pharmaceutically acceptable carriers and/or additives.

20 8. The use of at least one of the compounds as claimed in any of claims 1 to 6 for preparing a pharmaceutical composition for treating immunologically mediated inflammatory diseases.

25 9. A method for treating immunologically mediated inflammatory diseases, wherein an amount of a compound of the formula I as claimed in any of claims 1 to 6 sufficient to have immunomodulating action and/or to inhibit the release of cytokine is administered to a  
30 person in need of such a treatment.